

### (19) World Intellectual Property Organization International Bureau



### 

## (43) International Publication Date21 September 2000 (21.09.2000)

### **PCT**

# (10) International Publication Number WO 00/55364 A3

(51) International Patent Classification7:

C12Q 1/68

- (21) International Application Number: PCT/GB00/00916
- (22) International Filing Date: 10 March 2000 (10.03.2000)
- (25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

99301933.0

12 March 1999 (12.03.1999) EP

- (71) Applicant (for all designated States except US): AMER-SHAM PHARMACIA BIOTECH UK LIMITED [GB/GB]; Amersham Place, Little Chalfont, Buckinghamshire HP7 9NA (GB).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): REEVE, Michael, Alan [GB/GB]: 118 St Andrews Road, Henley-on-Thames, Oxfordshire RG9 1PL (GB). WORKMAN, Nicholas, Ian [GB/GB]; 4 Canford Court, Aylesbury, Buckinghamshire HP21 7NZ (GB). MARTIN-PARRAS, Luis [ES/GB]; 7 Gladstone Rise, High Wycombe, Buckinghamshire HP13 7NW (GB).

- (74) Agents: ROLLINS, Anthony, John et al.; Amersham Pharmacia Biotech UK Limited, Amersham Laboratories, White Lion Road, Amersham, Buckinghamshire HP7 9LL (GB).
- (81) Designated States (national): AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

#### Published:

- with international search report
- (88) Date of publication of the international search report: 11 October 2001

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: GENETIC ANALYSIS

(57) Abstract: A method is described for use in whole genome analysis. The method - termed inter-population perfectly matched duplex depletion - can overcome many of the limitations of current approaches based upon SNPs and linkage disequilibrium within isolated populations. Inter-population perfectly matched duplex depletion isolates a fragment (or fragments) containing differences between the "affected" and "unaffected" populations or cells. A convenient method - terminal restriction site profiling arrays (TRSPAs) - is described for the analysis of such fragments. A totally diagnostic internal control DNA is also described which allows both the extent and exact nature of any partial digestion to be unambiguously determined for inter-population perfectly matched duplex depletion or TRSPA restriction.